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INFORMATION DISCLOSURE STATEMENT BY APPLICANT (Use as many sheets as necessary)		Complete if Known	
		Application Number	10/529,511
		Filing Date	§371 of PCT/IL2003/000764
		First Named Inventor	Hermona Soreq
		Art Unit	
		Examiner Name	
Sheet 1	of 6	Attorney Docket Number	74136/JPW/JW

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T ²
RK	1 /	Abbott C. A., Mackness, M. I., Kumar, S., Olukoga, O., Gordon, C., Arrol, S., Bhatagar, D., Boulton, A. J. M., and Durrington, P. N. (1993) Relationship between serum butyrylcholinesterase activity, hypertriglyceridaemia and insulin sensitivity in diabetes mellitus. <i>Clin. Sci. (Lond)</i> 85: 77-81	
	2 /	Adkins, S., Gan, K.N., Mody, M., and La Du, B. N. (1993) Molecular basis for the polymorphic forms of human serum paraoxonase/arylesterase: glutamine or arginine at position 191, for the respective A or B allozymes. <i>Am. J Hum. Genet.</i> 52: 598-608	
	3 /	Akhmedova, S., Anisimov, S., Yakimovsky, A., and Schwartz, E. (1999) Gln → Arg 191 polymorphism of paraoxonase and Parkinson's disease. <i>Hum. Hered.</i> 49: 178-180	
	4	Aminoff, M. J. (2001) Parkinson's disease and other extrapyramidal disorders. In: Braunwald, E., et al. (eds) Harrison's principles of internal medicine. McGraw Hill, pp. 2399-2406	
	5	Bartels, C. F., Jensen, F. S., Lockridge, O., van der Spek, A. F., Rubinstein, H. M., Lubrano, T., and La Du, B. N. (1992) DNA mutation associated with the human butyrylcholinesterase K-variant and its linkage to the atypical variant mutation and other polymorphic sites. <i>Am. J. Hum. Genet.</i> 50: 1086-1103	
	6	Betarbet, R., Sherer, T. B., MacKenzie, G., Garcia-Osuna, M., Panov, A. V., and Greenamyre, J. T. (2000) Chronic systemic pesticide exposure reproduces features of Parkinson's disease. <i>Nature Neurosci.</i> 3: 1301-1306	
	7	Brindle, N., Song, Y., Rogaeva, E., Premkumar, S., Levesque, G., Yu, G., Ikeda, M., Nishimura, M., Paterson, A., Sorbi, S., Duara, R., Farrer, L., and St George-Hyslop, P. (1998) Analysis of the butyrylcholinesterase gene and nearby chromosome 3 markers in Alzheimer disease. <i>Hum. Mol. Genet.</i> 7: 933-935	
	8	Brophy, V. H., Jampsa, R. L., Clendenning, J. B., McKinstry, L. A., Jarvik, G. P., and Furlong, C. E. (2001) Effects of 5' regulatory-region polymorphisms on paraoxonase-gene (PON1) expression. <i>Am. J. Hum. Genet.</i> 68: 1428-1436	
	9	Brophy, V. H., Hastings, M. D., Clendenning, J. B., Richter, R. J., Jarvik, G. P., and Furlong, C. E. (2001) Polymorphisms in the human paraoxonase (PON1) promoter. <i>Pharmacogenetics</i> 11: 77-84	
✓	10	Burkhardt, C., Kelly, J. P., Lim, Y. H., Filley, C. M., and Parker, W. D. Jr. (1993) Neuroleptic medications inhibit complex I of the electron transport chain. <i>Ann. Neurol.</i> 33: 512-517	

Examiner Signature	<i>Hermona Soreq</i>	Date Considered	10/20/06
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Applicants: Hermona Soreq, et al.
 U.S. Serial No. 10/529,511
 Filed: as §371 national stage of
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 Exhibit A

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Sheet 2 of 6

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RK	11	Cassarino, D. S., Fall, C. P., Swerdlow, R. H., Smith, T. S., Halvorsen, E. M., Miller, S. W., Parks, J. P., Parker, W. D. Jr., and Bennett, J. P. Jr. (1997) Elevated reactive oxygen species and antioxidant enzyme activities in animal and cellular models of Parkinson's disease. <i>Biochim. Biophys. Acta</i> , 1362: 77-86	
	12	Costa, L. G., Cole, T. B., Jarvik, G. P., and Furlong, C. E. (2003) Functional genomic of the paraoxonase (PON1) polymorphisms: effects on pesticide sensitivity, cardiovascular disease, and drug metabolism. <i>Annu. Rev. Med.</i> 54: 371-392	
	13	Costa, L. G., Richter, R. J., Murphy, S. D., Omenn, G. S., Motulsky, A. G., and Furlong, C. E. (1987) Species differences in serum paraoxonase correlate with sensitivity to paraoxon toxicity. In: Costa, L. G. (eds.) <i>Toxicology of pesticides: experimental, clinical and regulatory perspectives</i> . Springer-Verlag, Heidelberg, pp. 263-266	
	14	Costa, L. G., Li, W. F., Richter, R. J., Shih, D. M., Lusia, A., and Furlong, C. E. (1999) The role of paraoxonase (PON1) in the detoxication of organophosphates and its human polymorphism. <i>Chem. Biol. Interact.</i> 119-120: 429-438	
	15	Dempster, A. P., Laird, N. M., and Rubin, D. B. (1977) Maximum Likelihood from Incomplete Data via the EM Algorithm. <i>J. Royal Statist. Soc. Ser. B.</i> 39	
	16	Furlong, C. E., Li, W. F., Costa, L. G., Richter, R. J., Shih, D. M., and Lusia, A. J. (1998) Genetically determined susceptibility to organophosphorus insecticides and nerve agents: developing a mouse model for the human PON1 polymorphism. <i>Neurotoxicology</i> 19: 645-650	
	17	Goldsmith, J. R., Herishanu, Y., Abarbanel, J. M., and Weinbaum, Z. (1990) Clustering of Parkinson's Disease Points to Environmental Etiology. <i>Arch. Environ. Health</i> 45: 88-94	
	18	Haley, R. W., Billecke, S., and La Du, B. N. (1999) Association of low PON1 type Q (type A) arylesterase activity with neurologic symptom complexes in Gulf War veterans. <i>Toxicol. Appl. Pharmacol.</i> 157: 227-233	
	19	Herishanu, Y. O., Goldsmith, J. R., Abarbanel, J. M., and Weinbaum, Z. (1989) Clustering of Parkinson's Disease in Southern Israel. <i>Can. J. Neurol. Sci.</i> 16: 402-405	
✓	20	Hodgson, E. and Lewy, P. E. (1996) Pesticides: An Important but Underused Model for the Environmental Health Sciences. <i>Environ. Health Perspect.</i> 104: 97-106	

Examiner Signature	<i>Kerame Koon</i>	Date Considered	10/20/06
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RK	21	Jenner, P. and Olanow, C. W. (1996) Oxidative stress and the pathogenesis of Parkinson's disease. <i>Neurology</i> 47(Suppl 3): S161-S170	
	22	Kitada, T., Asakawa, S., Hattori, N., Matsumine, H., Yamamura, Y., Minoshima, S., Yokochi, M., Mizuno, Y., and Shimizu, N. (1998) Mutation in the parkin gene cause autosomal recessive juvenile parkinsonism. <i>Nature</i> 392: 605-608	
	23	Kruger, R., Kuhn, W., Muller, T., Woitalla, D., Graeber, M., Kosel, S., Przuntek, H., Epplen, J. T., Schols, L., and Riess, O. (1998) Ala30Pro mutation in the gene encoding α -synuclein in Parkinson's disease. <i>Nat. Genet.</i> 18: 106-108	
	24	Lang, A. E. and Lozano, A. M. (1998) Parkinson's Disease. First of Two Parts. <i>N. Engl. J. Med.</i> 339: 1044-1053	
	25	Le Couteur, D. G., Muller, M., Yang, M. C., Mellick, G. D., and McLean, A. J. (2002) Age-environment and gene-environment interactions in the pathogenesis of Parkinson's disease. <i>Rev. Environ. Health</i> 17: 51-64	
	26	Lehmann, D. J., Nagy, Z., Litchfield, S., Borja, M. C., and Smith, A. D. (2000) Association of butyrylcholinesterase K variant with cholinesterase-positive neuritic plaques in the temporal cortex in late-onset Alzheimer's disease. <i>Hum. Genet.</i> 106: 447-452	
	27	Lehmann, D. J., Johnston, C., and Smith, A. D. (1997) Synergy between the genes for butyrylcholinesterase K variant and apolipoprotein E4 in late-onset confirmed Alzheimer's disease. <i>Hum. Mol. Genet.</i> 6: 1933-1936	
	28	Lockridge, O. and Masson, P. (2000) Pesticides and Susceptible Populations: People With Butyrylcholinesterase Genetic Variants May Be At Risk. <i>Genetic Neurotoxicology</i> 21: 113-126	
	29	Loewenstein-Lichtenstein, Y., Schwarz, M., Glick, D., Norgaard-Pedersen, B., Zakut, H., and Soreq, H. (1995) Genetic predisposition to adverse consequences of anti-cholinesterases in 'atypical' BCHE carriers. <i>Nat. Med.</i> 1: 1225-1226	
	30	Lucking, C. B., Durr, A., Bonifati, V., Vaughan, J., De Michele, G., Gasser, T., Harhangi, B. S., Meco, G., Deneffe, P., Wood, N. W., Agid, Y., and Brice, A. (2000) Association between early-onset Parkinson's disease and mutations in the parkin gene. French Parkinson's Disease Genetics Study Group. <i>N. Engl. J. Med.</i> 342: 1560-1567	

Examiner Signature	<i>Koranne Kozov</i>	Date Considered	10/20/06
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RK	31	Mackness, B., Durrington, P. N., and Mackness, M. I. (1998) Human Serum Paraoxonase. <i>Gen. Pharmacol.</i> 31: 329-336	
	32	Masson, P., Josse, D., Lockridge, O., Vigue, N., Taupin, C., and Buhler, C. (1998) Enzymes hydrolyzing organophosphates as potential catalytic scavengers against organophosphate poisoning. <i>J. Physiol. (Paris)</i> 92: 357-362	
	33	Menegon, A., Board, P. G., Blackburn, A. C., Mellick, G. D., and Le Couteur, D. G. (1998) Parkinson's disease, pesticides, and glutathione transferase polymorphisms. <i>Lancet</i> 352: 1344-1346	
	34	Nassar, B. A., Dunn, J., Title, L. M., O'Neill, B. J., Kirkland, S. A., Zayed, E., Bata, I. R., Cantrill, R. C., Johnstone, J., Dempsey, G. I., Tan, M. H., Breckenridge, W. C., and Johnstone, D. E. (1999) Relation of genetic polymorphisms of apolipoprotein E, angiotensin converting enzyme, apolipoprotein B-100, and glycoprotein IIIa and early-onset coronary heart disease. <i>Clin. Biochem.</i> 32: 275-282	
	35	Nassar, B. A., Darvesh, S., Bevin, L. D., Rockwood, K., Kirkland, S. A., O'Neill, B. J., Bata, I. R., Johnstone, D. E., and Title, L. M. (2002) Relation between butyrylcholinesterase K variant, paraoxonase I (PON1) Q and R and apolipoprotein E ε4 genes in early-onset coronary artery disease. <i>Clin. Biochem.</i> 5: 205-209	
	36	Parker, W. D. and Swerdlow, R. H. (1998) Mitochondrial dysfunction in idiopathic Parkinson disease. <i>Am. J. Hum. Genet.</i> 62: 758-762	
	37	Polymeropoulos, M. H., Lavedan, C., Leroy, E., Ide, S. E., Dehejia, A., Dutra, A., Pike, B., Root, H., Rubenstein, J., Boyer, R., Stenroos, E. S., Chandrasekharappa, S., Athanassiadou, A., Papapetropoulos, T., Johnson, W. G., Lazzarini, A. M., Duvoisin, R. C., Di Iorio, G., Golbe, L. I., and Nussbaum, R. L. (1997) Mutation in the α-synuclein gene identified in families with Parkinson's disease. <i>Science</i> 276: 2045-2047	
	38	Premkumar, D. R., Cohen, D. L., Hedera, P., Friedland, R. P., and Kalaria, R. N. (1996) Apolipoprotein E-ε4 alleles in cerebral amyloid angiopathy and cerebrovascular pathology associated with Alzheimer's disease. <i>Am. J. Pathol.</i> 148: 2083-2095	
	39	Spillantini, M. G., Schmidt, M. L., Lee, V. M.-Y., Trojanowski, J. Q., Jakes, R., and Goedert, M. (1997) α-Synuclein in Lewy Bodies. <i>Nature</i> 388: 839-40	
✓	40	Suehiro, T., Nakamura, T., Inoue, M., Shiinoki, T., Ikeda, Y., Kumon, Y., Shindo, M., Tanaka, H., and Hashimoto, K. (2000) A polymorphism upstream from the human paraoxonase (PON1) gene and its association with PON1 expression. <i>Atherosclerosis</i> 150: 295-298	

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RVL	41	Sveinbjörnsdóttir, S., Hicks, A. A., Jónsson, T., Péturason, H., Guðmundsson, G., Grigge, M. L., Kong, A., Gulcher, J. R., and Stefánsson, K. (2000) Familial Aggregation of Parkinson's Disease in Iceland. <i>N. Engl. J. Med.</i> 343: 1765-1770	
	42	Tanner, C. M., Ottman, R., Ellenberg, J. H., Goldman, S. M., Mayeux, R., Chan, P., and Langston, J. W. (1997) Parkinson's Disease (PD) Concordance in Elderly Male Monozygotic (MZ) and Dizygotic (DZ) Twins. <i>Neurology</i> 48(Suppl): A333	
	43	Taylor, M. C., Le Couteur, D. G., Mellick, G. D., and Board, P. G. (2000) Paraoxonase polymorphisms, pesticide exposure and Parkinson's disease in a Caucasian population. <i>J. Neural Transm.</i> 107: 979-983	
	44	Vays, I., Heikkila, R. E., and Nicklas, W. J. (1986) Studies on the neurotoxicity of 1-methyl-4-phenyl-1,2,3,6-tetrahydropyridine: inhibition of NAD-linked substrate oxidation by its metabolite, 1-methyl-4-phenylpyridinium. <i>J. Neurochem.</i> 46: 1501-1507	
	45	Vingerhoets, F. J. G., Snow, B. J., Tetud, J. W., Langston, J. W., Schulzer, M., and Calne, D. B. (1994) Positron emission tomographic evidence for progression of human MPTP-induced dopaminergic lesions. <i>Ann. Neurol.</i> 36: 765-770	
	46	Wang, J. and Liu, Z. (2000) No association between paraoxonase 1 (PON1) gene polymorphisms and susceptibility to Parkinson's disease in a Chinese population. <i>Mov. Disord.</i> 15: 1265-1267	
	47	Wooten, G. F., Currie, L. J., Bennett, J. P., Harrison, M. B., Trugman, J. M., and Parker, W. D. Jr. (1997) Maternal inheritance in Parkinson's disease. <i>Ann. Neurol.</i> 41: 265-268	
	48	Poewe, W. H. and Wenning, G. K. (1996) The natural history of Parkinson's disease. <i>Neurology</i> 47(Suppl 3): S146-S152	
	49	International Preliminary Examination Report issued by the International Preliminary Examination Authority (IPEA/EP) on December 27, 2004 in connection with International Application No. PCT/IL2003/000764	
	50	International Search Report issued by the International Searching Authority (ISA/EP) on April 6, 2004 in connection with International Application No. PCT/IL2003/000764	

Examiner
Signature

Cheranne Korman

Date

Considered

10/20/06

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PK	51	Akhmedova, S. N., Yakimovsky, A. K., and Schwartz, E. I. (2001) Paraoxonase I Met-Leu 54 Polymorphism Is Associated With Parkinson's Disease. <i>J. Neurol. Sci.</i> 184: 179-182	
	52	Shapira, M., Tur-Kaspa, I., Bosgraaf, L., Livni, N., Grant, A. D., Grisaru, D., Komer, M., Ebstein, R. P., and Soreq, H. (2000) A transcription-activating polymorphism in the AChE promoter associated with acute sensitivity to anti-acetylcholinesterases. <i>Hum. Mol. Genet.</i> 9: 1273-1281	
	53	Kondo, I. and Yamamoto, M. (1998) Genetic Polymorphism Of Paraoxonase I (PON1) And Susceptibility To Parkinson's Disease. <i>Brain Res.</i> , 806: 271-273	
	54	Bartels, C. F., Zelinski, T., and Lockridge, O. (1993) Mutation at Codon 322 in the Human Acetylcholinesterase (ACHE) Gene Accounts for YT Blood Group Polymorphism. <i>Amer. J. Hum. Genet.</i> 52: 928-936	
	55	Carmine, A., Buervenich, S., Sydow, O., Anvret, M., and Olson, L. (2002) Further Evidence for an Association of the Paraoxonase I (PON1) Met-54 Allele with Parkinson's Disease. <i>Movement Disord.</i> 17: 764-766	
	56	Kaufer, D. and Soreq, H. (1999) Tracking Cholinergic Pathways from Psychological and Chemical Stressors to Variable Neurodeterioration Paradigms. <i>Curr. Opin. Neurol.</i> 12: 739-743	

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